& Fin Delwal

98817

Access DB#\_\_\_\_

# SEARCH REQUEST FORM

Scientific and Technical Inf rmation Center

2819		Examiner #: 1/1469 Date: 1/15/03  Serial Number: 10/046120  Ilts Format Preferred (circle): PAPER DISK E-MAIL  e searches in order of need.
Include the elected species or structures, key	ywords, synonyms, acron nat may have a special me	**************************************
Title of Invention: Treatm	ent o sle	ep posturbance
Inventors (please provide full names):	kack eta	ep bisturbance
		*
Earliest Priority Filing Date:	9/01	·
,	• •	parent, child, divisional, or issued patent numbers) along with the
appropriate serial number.	an permen myermanen (	
1) Please search	clais ,	1 — 13
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		Jan Delaval Reference Librarian Biotechnology & Chemical Library CM1 1E07 ~ 703-308-4498 jan.delaval@uspto.gov
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Searcher:	NA Sequence (#)	STN
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Searcher Location:	Structure (#)	Questel/Orbit
Date Searcher Picked Up:	Bibliographic	Dr.Link
Date Completed:	Litigation	Lexis/Nexis
Searcher Prep & Review Time:	Fulltext	Sequence Systems
Clerical Prep Time:	Patent Family	WWW/Internet
Online Time: + 44	Other	Other (specify)

PTO-1590 (8-01)



# STIC Search Report Biotech-Chem Library

## STIC Database Tracking Number: 98817

TO: Jennifer Kim

Location: 2d17(12b19)

Wednesday, July 16, 2003

Art Unit: 1617 Phone: 308-2232

Serial Number: 10 / 046727

From: Jan Delaval

**Location: Biotech-Chem Library** 

CM1-1E07

Phone: 308-4498

jan.delaval@uspto.gov

### Search Notes

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# STIC SEARCH RESULTS

# **Biotech-Chem Library**

Questions about the scope or the results of the search? Contact the searcher or contact:

Mary Hale, Information Branch Supervisor 308-4258, CM1-1E01

VO	untary Results reedback rum
<b>A</b>	I am an examiner in Workgroup: Example: 1610
>	Relevant prior art found, search results used as follows:
	☐ 102 rejection
	☐ 103 rejection
	☐ Cited as being of interest.
	Helped examiner better understand the invention.
	Helped examiner better understand the state of the art in their technology.
	Types of relevant prior art found:
	☐ Foreign Patent(s)
	<ul> <li>Non-Patent Literature (journal articles, conference proceedings, new product announcements etc.)</li> </ul>
>	Relevant prior art not found:
	Results verified the lack of relevant prior art (helped determine patentability).
	Results were not useful in determining patentability or understanding the invention.
Co	omments:

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Biotechnology & Chemical Library
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This file contains CAS Registry Numbers for easy and accurate substance identification.

#### => d all hitstr tot 198

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L98 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2003 ACS
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AN 2003:117637 HCAPLUS

DN 138:158841

TI Sucralose formulations to mask unpleasant tastes

IN Jaeger, David; Dickerson, Jay; Mogavero, Annabelle

PA Wyeth, John, and Brother Ltd., USA

SO PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K031-715

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

```
DATE
                                          APPLICATION NO.
    PATENT NO.
                     KIND
                     ----
                           _____
                                          ______
                                                           _____
    WO 2003011306
                      Α1
                           20030213
                                          WO 2002-US24298 20020731
PI
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
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            TJ, TM
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            PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
            NE, SN, TD, TG
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PRAI US 2001-308912P P 20010731

AB The present invention is directed to a pharmaceutically acceptable taste masking liq. excipient base for administration of a relatively large amt. of unpleasant tasting medicines. More particularly, the enhanced sweetness and taste masking effect are produced by the addn. of sucralose to the excipient base with maintenance of a pH 2-5. The invention is further directed to medicinal compns. comprising such a liq. excipient base and unpleasant tasting medicines. Still further, the invention is directed to a method for taste masking unpleasant tasting medicines through their incorporation into the claimed liq. excipient bases. A

cough syrup , contained quaifenesin 2.00, pseudoephedrine-HCl 0.60, dextromethorphan-HBr 0.20, PEG-1450 20.00, glycerin 6.00, high-fructose corn syrup (55%) 45.00, anhyd. citric acid 1.60, and 70% sorbitol 5.00, sodium citrate 0.20, sodium benzoate 0.30, Red No. 40 dye 0.05, golden punch flavor 0.10, sucralose 0.20, and water to 100.0%. ST sucralose formulation unpleasant taste masking ΙT Anti-inflammatory agents (nonsteroidal; sucralose formulations to mask unpleasant tastes) ΙT Analgesics Antibiotics Antihistamines Antitumor agents Antitussives Bitterness Decongestants Expectorants Human (sucralose formulations to mask unpleasant tastes) IT Drug delivery systems (syrups, cough; sucralose formulations to mask unpleasant tastes) 52-28-8, Codeine phosphate 58-73-1, Diphenhydramine IT 59-42-7, Phenylephrine 77-22-5, Caramiphen 59-33-6, Pyrilamine 86-22-6, Brompheniramine 91-81-6, Tripelenamine 93-14-1, Guaifenesin 113-92-8, Chlorpheniramine maleate 103-90-2, Acetaminophen 125-69-9, Dextromethorphan hydrobromide Bromodiphenhydramine 132-21-8, DexBrompheniramine 345-78-8, Pseudoephedrine Cyproheptadine 486-12-4, Triprolidine 562-10-7, Doxylamine hydrochloride 980-71-2, Brompheniramine maleate 1321-14-8, Potassium Phenindamine quaicolsulfonate 1420-53-7, Codeine sulfate 2451-01-6, Terpin hydrate 3964-81-6, Azatadine 7681-11-0, Potassium iodide (KI), biological 7778-49-6, Potassium citrate 14838-15-4, Phenylpropanolamine 22071-15-4, Ketoprofen 15687-27-1, Ibuprofen 25523-97-1, DexChlorpheniramine 56038-13-2, 22204-53-1, Naproxen Sucralose RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (sucralose formulations to mask unpleasant tastes) THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Blase; US 5409907 A 1995 HCAPLUS (2) Cherukuri; US 5013716 A 1991 HCAPLUS IT 58-73-1, Diphenhydramine 15687-27-1, Ibuprofen RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (sucralose formulations to mask unpleasant tastes) RN 58-73-1 HCAPLUS Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl- (9CI) (CA INDEX NAME) CNPh2CH-O-CH2-CH2-NMe2 15687-27-1 HCAPLUS RN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) CN NAME)

```
HO<sub>2</sub>C-CH Me
```

```
ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2003 ACS
L98
AN
     2002:555345 HCAPLUS
DN
     137:103930
ΤI
     Ibuprofen-diphenhydramine composition and method for
     the treatment of pain-associated sleep disturbances
     Cooper, Stephen Allen; Doyle, Geraldine Ann;
IN
     Cook, Graham David; Koch, Todd Shawn; Fort, James
     Joseph; Bianco, Giustino; Giamalva, David Hugh
     Wyeth, John, and Brother Ltd., USA
PA
SO
     PCT Int. Appl., 75 pp.
     CODEN: PIXXD2
DT
     Patent
     English
LA
     ICM A61K031-00
IC
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 63
FAN.CNT 1
     PATENT NO.
                      KIND
                           DATE
                                           APPLICATION NO.
                                                            DATE
                                                           _____
                                           -----
                            _____
     WO 2002056877 · A1
                            20020725
                                           WO 2002-US3855
                                                            20020118 <--
PΙ
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ,
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
             CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     US 2002177626
                      A1
                            20021128
                                           US 2002-46727
                                                            20020117 <--
PRAI US 2001-262387P
                     Р
                            20010119
     The invention provides a compn. for treating pain-assocd. sleep
     disturbances, esp. shortened sleep duration, comprising
     ibuprofen and diphenhydramine. The compn. is further
     prepd. as a bilayer tablet or caplet, or alternatively as a soft gelatin
     capsule compn., to prevent interaction between the active ingredients.
     ibuprofen diphenhydramine pharmaceutical pain assocd
ST
     sleep disturbance treatment
IT
     Drug delivery systems
        (caplets, bilayer; ibuprofen-diphenhydramine compn.
        and method for treatment of pain-assocd. sleep disturbances)
IT
     Drug delivery systems
        (capsules; ibuprofen-diphenhydramine compn. and
        method for treatment of pain-assocd. sleep disturbances)
IT
     Sleep
        (disorder; ibuprofen-diphenhydramine
        compn. and method for treatment of pain-assocd. sleep
        disturbances)
IT
     Analgesics
     Drug delivery systems
     Drug interactions
     Human
```

(CA INDEX

```
Pain
        (ibuprofen-diphenhydramine compn. and method for
        treatment of pain-assocd. sleep disturbances)
TΤ
     Polyoxyalkylenes, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ibuprofen-diphenhydramine compn. and method for
        treatment of pain-assocd. sleep disturbances)
TΤ
     Surgery
        (oral; ibuprofen-diphenhydramine compn. and method
        for treatment of pain-assocd. sleep disturbances)
     Drug delivery systems
ፐጥ
        (tablets, bilayer; ibuprofen-diphenhydramine compn.
        and method for treatment of pain-assocd. sleep disturbances)
     58-73-1, Diphenhydramine 147-24-0,
ΤТ
     Diphenhydramine hydrochloride 15687-27-1,
     Ibuprofen 88637-37-0, Diphenhydramine
     citrate
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (ibuprofen-diphenhydramine compn. and method for
        treatment of pain-assocd. sleep disturbances)
ΙT
     25322-68-3, Polyethylene glycol
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ibuprofen-diphenhydramine compn. and method for
        treatment of pain-assocd. sleep disturbances)
RE.CNT
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
(1) Cevc Gregor; EP 0475160 A 1992 HCAPLUS
(2) Charles, F; WO 9729735 A 1997 HCAPLUS
(3) Drizen, A; US 6063405 A 2000 HCAPLUS
(4) Lam Pharmaceuticals Inc; WO 9711681 A 1997 HCAPLUS
(5) Sunshine, A; US 4522826 A 1985 HCAPLUS
(6) Sunshine, A; US 4985459 A 1991 HCAPLUS
IT
     58-73-1, Diphenhydramine 147-24-0,
     Diphenhydramine hydrochloride 15687-27-1,
     Ibuprofen 88637-37-0, Diphenhydramine
     citrate
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (ibuprofen-diphenhydramine compn. and method for
        treatment of pain-assocd. sleep disturbances)
RN
     58-73-1 HCAPLUS
     Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl- (9CI)
CN
                                                           (CA INDEX NAME)
Ph2CH-O-CH2-CH2-NMe2
RN
     147-24-0 HCAPLUS
     Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl-, hydrochloride (9CI)
CN
                                                                           (CA
     INDEX NAME)
Ph2CH-O-CH2-CH2-NMe2
           HCl
RN
     15687-27-1 HCAPLUS
```

Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI)

CN

NAME)

RN 88637-37-0 HCAPLUS

CN Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl-, 2-hydroxy-1,2,3-propanetricarboxylate (1:1) (9CI) (CA INDEX NAME)

CM ·1

CRN 77-92-9 CMF C6 H8 O7

CM 2

CRN 58-73-1 CMF C17 H21 N O

Ph2CH-O-CH2-CH2-NMe2

IT 25322-68-3, Polyethylene glycol

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ibuprofen-diphenhydramine compn. and method for treatment of pain-assocd. sleep disturbances)

RN 25322-68-3 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy- (9CI) (CA INDEX NAME)

$$HO = CH_2 - CH_2 - O = H$$

L98 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:157572 HCAPLUS

DN 136:189396

TI **Ibuprofen** solutions for **capsule**-filling and **capsule** preparations

IN Kato, Soichiro; Tsumori, Katsuyuki; Shinoda, Yasuo; Inaqi, Toshio

PA Kowa Company, Ltd., Japan

SO PCT Int. Appl., 18 pp. CODEN: PIXXD2

DT Patent

```
LA
     Japanese'
IC
     ICM A61K031-192
         A61K009-08; A61K009-48; A61K047-34; A61K047-10; A61K047-06;
         A61K047-08
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                                                           _____
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                           -----
                                           -----
                            20020228
                                          WO 2001-JP7223 20010823 <--
     WO 2002015900
                      A1
ΡI
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             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
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             US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                                           20010823 <--
     AU 2001080124
                      Α5
                            20020304
                                          AU 2001-80124
                                                          20010823 <--
     EP 1321140
                            20030625
                                           EP 2001-958434
                       A1
             AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
         R:
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
PRAI JP 2000-255827
                      Α
                            20000825
                                      <--
     WO 2001-JP7223
                      W
                            20010823
     Capsule prepns. are produced by using solns. for capsule
     -filling which contain ibuprofen, polyethylene
     glycol, water and terpenes (menthol, limonene, borneol,
     dl-camphor, mentha oil, etc.) optionally together with one or more drugs
     selected from among antipyretic analgesics, antihistamines, antitussives,
     expectorants, sympathetic stimulants, analeptics, hypnotic sedatives and
     anti-inflammatory agents, solubilizers, thickeners, pH controlling agents,
     colorants, etc. Thus, it is attempted to relieve the bitterness of
     ibuprofen and establish an immediate action at the same time.
ST
     capsule ibuprofen PEG terpene
ΙT
     Antipyretics
        (analgesic; capsules contg. ibuprofen and
        PEG and terpenes and other drugs)
ΙT
     Analgesics
        (antipyretic; capsules contg. ibuprofen and
        PEG and terpenes and other drugs)
IT
     Anti-inflammatory agents
     Antihistamines
     Antitussives
     Expectorants
     Hypnotics and Sedatives
     Nervous system stimulants
        (capsules contg. ibuprofen and PEG and
        terpenes and other drugs)
     Polyoxyalkylenes, biological studies
IT
     Terpenes, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (capsules contg. ibuprofen and PEG and
        terpenes and other drugs)
IT
     Drug delivery systems
        (capsules; capsules contg. ibuprofen and
        PEG and terpenes and other drugs)
IT
     Essential oils
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (mint, Mentha; capsules contg. ibuprofen and
        PEG and terpenes and other drugs)
IT
     Nerve
        (sympathetic, stimulants; capsules contg. ibuprofen
        and PEG and terpenes and other drugs)
```

```
ΙT
     58-08-2, Caffeine, biological studies
                                             76-22-2, dl-Camphor
                                                                    89-78-1.
               93-14-1, Guaifenesin 113-92-8, Chlorpheniramine maleate
     125-69-9, Dextromethorphan hydrobromide
                                              138-86-3, Limonene
     147-24-0, Diphenhydramine hydrochloride
                         912-60-7, Noscapine hydrochloride 15687-27-1
     507-70-0, Borneol
                   18760-80-0, dl-Methylephedrine hydrochloride
     , Ibuprofen
     24204-13-5, Dihydrocodeine phosphate 25322-68-3,
     Polyethylene glycol
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (capsules contg. ibuprofen and PEG and
        terpenes and other drugs)
RE.CNT 21
              THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Consumer Health S A; ZA 9706189 A 1998 HCAPLUS
(2) Consumer Health S A; ZA 9706189 A 1998 HCAPLUS
(3) Consumer Health S A; AU 9734442 A 1998 HCAPLUS
(4) Consumer Health S A; AU 9734442 A 1998 HCAPLUS
(5) Consumer Health S A; WO 9802182 A1 1998 HCAPLUS
(6) Consumer Health S A; WO 9802182 A1 1998 HCAPLUS
(7) R P Scherer Corporation; JP 01502185 A 1988
(8) R P Scherer Corporation; EP 293406 A1 1988 HCAPLUS
(9) R P Scherer Corporation; US 5071643 1988 HCAPLUS
(10) R P Scherer Corporation; US 5360615 1988 HCAPLUS
(11) R P Scherer Corporation; ZA 8708723 A 1988 HCAPLUS
(12) R P Scherer Corporation; AU 8781573 A 1988 HCAPLUS
(13) R P Scherer Corporation; WO 8802625 A1 1988 HCAPLUS
(14) Taisho Pharmaceutical Co Ltd; JP 08333246 A 1996 HCAPLUS
(15) Taisho Pharmaceutical Co Ltd; US 5827852 A 1996 HCAPLUS
(16) The Procter & Gamble Company; JP 08509725 A 1994
(17) The Procter & Gamble Company; US 5827852 A 1994 HCAPLUS
(18) The Procter & Gamble Company; EP 695175 Al 1994 HCAPLUS
(19) The Procter & Gamble Company; WO 9425009 A1 1994 HCAPLUS
(20) The Procter & Gamble Company; NO 9504146 A 1994 HCAPLUS
(21) The Procter & Gamble Company; FI 9505144 A 1994 HCAPLUS
IT
     147-24-0, Diphenhydramine hydrochloride
     15687-27-1, Ibuprofen 25322-68-3,
     Polyethylene glycol
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (capsules contg. ibuprofen and PEG and
        terpenes and other drugs)
RN
     147-24-0 HCAPLUS
CN
     Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl-, hydrochloride (9CI)
                                                                          (CA
     INDEX NAME)
```

 $Ph_2CH-O-CH_2-CH_2-NMe_2$ 

#### HC1

RN 15687-27-1 HCAPLUS
CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) (CA INDEX NAME)

RN 25322-68-3 HCAPLUS

Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy- (9CI) (CA INDEX CN

$$HO \longrightarrow CH_2 - CH_2 - O \longrightarrow n$$

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L98
    ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2003 ACS
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ΑN 1999:27707 HCAPLUS

DN 130:86181

Pharmaceutical formulations containing ibuprofen and ΤI diphenhydramine analgesics

IN Hough, Douglas R.; Nelson, Edward B.; Raffa, Robert B.

PΑ McNeil-PPC, USA

SO PCT Int. Appl., 15 pp. CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K031-19

ICS A61K031-135; A61K031-19; A61K031-135

CC 63-6 (Pharmaceuticals)

Section cross-reference(s): 1

FAN.		1	ÇIO:	22-I	erer,	ence	(5).	1										
	PAT	CENT 1			KI		DATE					CATI			DATE			
ΡI	WO	9858					1998	1230							1997	0623	<	
		W:	AL,	AM,	AT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,
			DK,	EE,	ES,	FΙ,	GB,	GE,	HU,	IL,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	ΚZ,	LC,
		•	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,
			RO,	RU,	SD,	SE,	SG,	SI,	SK,	ТJ,	TM,	TR,	TT,	UA,	UG,	UZ,	VN,	AM,
			ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM								
		RW:					SD,	-	-							-	-	-
							LU,			PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,
							SN,						•					
		9743									J 199	97-43	3256		1997	0623	<	
PRAI		1997																
AB	AB Disclosed are compns. comprising ibuprofen (I) and																	
		phenh														. Wi	nen	
		prof																
	their pharmacol. effects are superadditive. ED50 of I and II.HCl																	
	decreased from 36.5 and 8.0 mg/kg, resp. to 2.4 mg/kg orally each, when																	
													noli	ne b	romi	de in	nduce	ed:
	combined and none of the mice tested in acetylcholine bromide induced abdominal constriction assay showed writhing.																	
	•			٠,	_				_	• • •								

pharmaceutical formulation ibuprofen diphenhydramine STsynergism analgesic

ITAlcohols, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (C16-18; pharmaceutical formulations contg. ibuprofen and diphenhydramine analgesics)

```
Quaternary ammonium compounds, biological studies
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (alkylbenzyldimethyl, chlorides; pharmaceutical formulations contg.
        ibuprofen and diphenhydramine analgesics)
     Polyoxyalkylenes, biological studies
ΙT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (alkylene derivs.; pharmaceutical formulations contg. ibuprofen
        and diphenhydramine analgesics)
     Emulsifying agents
IΤ
        (anionic; pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
     Fatty acids, biological studies
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (esters, with polyoxyethylene sorbitan; pharmaceutical
        formulations contg. ibuprofen and diphenhydramine
        analgesics)
IT
     Castor oil
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ethoxylated; pharmaceutical formulations contg. ibuprofen
        and diphenhydramine analgesics)
     Emulsifying agents
ΙT
        (nonionic; pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
ΙT
     Acacia
     Antihistamines
        (pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
IT
     Lanolin
     Lecithins
     Paraffin oils
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
ΙT
     Analgesics
        (synergistic; pharmaceutical formulations contg. ibuprofen
        and diphenhydramine analgesics)
IT
     58-73-1, Diphenhydramine 15687-27-1,
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
        (pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
                                                      57-55-6, 1,2-Propanediol,
ΙT
     57-11-4, Octadecanoic acid, biological studies
                          57-88-5, Cholest-5-en-3-ol (3.beta.)-, biological
     biological studies
               102-71-6, Triethanolamine, biological studies
                                                                111-42-2,
                                                                      120-51-4,
     biological studies
                          112-80-1, Oleic acid, biological studies
     Benzylbenzoate
                      121-54-0
                                 141-43-5, biological studies
                                                                151-21-3,
     Sodium lauryl sulfate, biological studies
                                                577-11-7, Docusate sodium
                                        9002-89-5
                                                     9004-64-2, Hydroxypropyl
     7558-79-4
                 8044-71-1, Cetrimide
                 9004-67-5, Methylcellulose
                                              9004-99-3,
                                9005-32-7, Alginic acid
                                                           9005-63-4D,
     Polyoxyethylene stearate
     Polyoxyethylene sorbitan, esters with fatty acids
                                                          12441-09-7D,
     Sorbitan, esters
                        12619-70-4, Cyclodextrin 25322-68-3D,
                        25496-72-4, Glyceryl monooleate
                                                           31566-31-1, Glyceryl
     alkylene derivs.
                                                106392-12-5, Poloxamer
     monostearate
                    36653-82-4, Cetyl alcohol
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical formulations contg. ibuprofen and
        diphenhydramine analgesics)
              THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Motola, S; US 5024997 A 1991 HCAPLUS
```

- (2) Richardson Vicks Inc; US 4522826 1985 HCAPLUS
- (3) Richardson Vicks Inc; WO 8503443 A 1985 HCAPLUS
- IT 58-73-1, Diphenhydramine 15687-27-1,

#### Ibuprofen

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical formulations contg. ibuprofen and diphenhydramine analgesics)

RN 58-73-1 HCAPLUS

CN Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl- (9CI) (CA INDEX NAME)

Ph2CH-O-CH2-CH2-NMe2

RN 15687-27-1 HCAPLUS

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI) (CA INDEX NAME)

IT 25322-68-3D, alkylene derivs.

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical formulations contg. ibuprofen and diphenhydramine analgesics)

RN 25322-68-3 HCAPLUS

CN Poly(oxy-1,2-ethanediyl), .alpha.-hydro-.omega.-hydroxy- (9CI) (CA INDEX NAME)

$$HO - \begin{bmatrix} CH_2 - CH_2 - O \end{bmatrix}_n H$$

L98 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2003 ACS

AN 1991:589755 HCAPLUS

DN 115:189755

TI Pharmaceutical composition and methods for treating the symptoms of overindulgence .

IN Goldman, William J.; Gates, Thomas N.

PA McNeil-PPC, Inc., USA

SO Eur. Pat. Appl., 8 pp. CODEN: EPXXDW

DT Patent

LA English

IC ICM A61K031-415

ICS A61K031-34; A61K031-165; A61K031-19; A61K031-44

ICI A61K031-415, A61K031-19, A61K031-165; A61K031-34, A61K031-165;
 A61K031-165, A61K031-135; A61K031-19, A61K031-135; A61K031-44, A61K031-19,
 A61K031-165

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

	PAT	ENT NO.	KIND	DATE	APPLICATION NO. DATE	
PI		426479 426479	A1 B1	19910508 19940216	EP 1990-311995 1990	1101
		R: AT, BE,	CH, DE	, ES, FR,	GB, IT, LI	
	IN	171746	À	19921226	IN 1990-CA907 1990	1029
	CA	2028746	AA	19910503	CA 1990-2028746 1990	1031
	CA	2028746	С	19951226		
	ΑU	9065689	A1	19910509	AU 1990-65689 1990	1031
	AU	646230	B2	19940217		
	ZA	9008775	Α	19920729	ZA 1990-8775 · 1990	1101
	AT	101515	E	19940315	AT 1990-311995 1990	1101
	ES	2057439	Т3	19941016	ES 1990-311995 1990	1101
	JP	03206052	A2 ·	19910909	JP 1990-298713 1990	1102
	US	5204118	A	19930420	00 200 0 000 000	0429
	IN	176327	Α.	19960427	IN 1992-CA757 1992	1016
	US	5417980	A	19950523	US 1994-268865 1994	0629
PRA	I US	1989-430837	A	19891102		
	IN	1990-CA907	A1	19901029		
	ΕP	1990-311995	A	19901101	•	
	ΰS	1992-876824	ĀЗ	19920429		
	US	1993-46534	В1	19930413		
ND	Dha		aamnna	which can	rolions the symptoms of on	arina

AB Pharmaceutical compns. which can relieve the symptoms of overindulgence, such as headache and acid indigestion, comprise (1) acetaminophen or a nonsteroidal antiinflammatory drug and (2) histamine H1 or H2 receptor blocker, proton pump inhibitor, or a combination thereof. The compn. can be in the form of tablets, capsules, or liqs. Thus, a tablet contained acetaminophen 500, cimetidine 150 mg, and other auxiliary agents and coloring agents.

ST overeating symptom antiinflammatory antihistaminic tablet; acetaminophen cimetidine tablet overeating symptom

IT Headache

(from overeating, treatment of, inflammation inhibitor-antihistaminic combinations for)

IT Inflammation inhibitors

(mixts. with antihistaminics, headache and indigestion from overeating treatment with)

IT Antacids and Antiflatulents

(mixts. with inflammation inhibitors, headache and indigestion from overeating treatment with)

IT Antihistaminics

(H1, mixts. with inflammation inhibitors, headache and indigestion from overeating treatment with)

IT Antihistaminics

(H2, mixts. with inflammation inhibitors, headache and indigestion from overeating treatment with)

IT Digestive tract

(disease, indigestion, from overeating, treatment of, inflammation inhibitor-antihistaminic combinations for)

IT Appetite

(disorder, hyperphagia, discomfort from, treatment of, inflammation inhibitor-antihistaminic combinations for)

IT Pharmaceutical dosage forms

(liqs., oral, of anti-inflammatory and antihistaminic combinations, for treatment of headache and indigestion from overeating)

IT Pharmaceutical dosage forms

(tablets, of anti-inflammatory and antihistaminic combinations, for treatment of headache and indigestion from overeating)

IT 53-86-1D, mixts. with antihistaminics 58-73-1D, mixts. with anti-inflammatory agent 59-33-6D, Pyrilamine, mixts. with anti-inflammatory agent 60-87-7D, Promethazine, mixts. with anti-inflammatory agent 61-68-7D, mixts. with antihistaminics 68-88-2D, Hydroxyzine, mixts. with anti-inflammatory agent 79-09-4D

82-92-8D, Propanoic acid, derivs., mixts. with antihistaminics Cyclizine, mixts. with anti-inflammatory agent 91-40-7D, Fenamic acid, derivs., mixts. with antihistaminics 91-81-6D, Tripelenamine, mixts. 103-90-2D, Acetaminophen, mixts. with with anti-inflammatory agent 113-92-8D, Chlorpheniramine, mixts. with antihistaminics 486-16-8D, Carbinoxamine, mixts. with anti-inflammatory agent 523-87-5D, mixts. with anti-inflammatory agent anti-inflammatory agent 644-62-2D, 569-65-3D, Meclizine, mixts. with anti-inflammatory agent 15687-27-1D, Ibuprofen, mixts. with mixts. with antihistaminics 22204-53-1D, 22071-15-4D, mixts. with antihistaminics antihistaminics mixts. with antihistaminics 26171-23-3D, Tolmetin, mixts. with 29679-58-1D, Fenoprofen, mixts. with antihistaminics antihistaminics 32536-43-9D, Indole acetic acid, derivs., mixts. with antihistaminics 38194-50-2D, 36322-90-4D, Piroxicam, mixts. with antihistaminics 51481-61-9D, Cimetidine, mixts. Sulindac, mixts. with antihistaminics 66357-35-5D, Ranitidine, mixts. with with anti-inflammatory agent 73590-58-6D, mixts. with anti-inflammatory agent anti-inflammatory agent 136584-95-7 76824-35-6D, mixts. with anti-inflammatory agent 136584-99-1 136585-00-7 136584-97-9 136584-98-0 136584-96-8 136585-04-1 136585-05-2 136585-01-8 136585-02-9 136585-03-0 136585-06-3 136585-07-4 **136585-08-5** RL: BIOL (Biological study) (headache and indigestion from overeating treatment with) 136585-08-5

ΙT

RL: BIOL (Biological study)

(headache and indigestion from overeating treatment with)

136585-08-5 HCAPLUS RN

CN Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, mixt. with 2-(diphenylmethoxy)-N, N-dimethylethanamine (9CI) (CA INDEX NAME)

CM

15687-27-1 CRN CMF C13 H18 O2

· CM 2

58-73-1 CRN CMF C17 H21 N O

Ph2CH-O-CH2-CH2-NMe2

=> sel hit rn 198 E7 THROUGH E12 ASSIGNED

=> fil reg;s e7-e12 FILE 'REGISTRY' ENTERED AT 17:09:58 ON 16 JUL 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 15 JUL 2003 HIGHEST RN 548735-19-9 DICTIONARY FILE UPDATES: 15 JUL 2003 HIGHEST RN 548735-19-9

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> d ide can tot

CM

ANSWER 1 OF 6 REGISTRY COPYRIGHT 2003 ACS RN 136585-08-5 REGISTRY Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)-, mixt. with CN 2-(diphenylmethoxy)-N, N-dimethylethanamine (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl-, mixt. contg. (9CI) C17 H21 N O . C13 H18 O2 MF CI MXS SR CA LC STN Files: CA, CAPLUS, USPATFULL

CRN 15687-27-1 CMF C13 H18 O2

1

CM 2

CRN 58-73-1 CMF C17 H21 N O

Ph2CH-O-CH2-CH2-NMe2

1 REFERENCES IN FILE CA (1957 TO DATE)
1 REFERENCES IN FILE CAPLUS (1957 TO DATE)

REFERENCE 1: 115:189755

L99 ANSWER 2 OF 6 REGISTRY COPYRIGHT 2003 ACS

RN **88637-37-0** REGISTRY

CN Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl-, 2-hydroxy-1,2,3-propanetricarboxylate (1:1) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Diphenhydramine citrate

MF C17 H21 N O . C6 H8 O7

CI COM

SR Commission of European Communities

LC STN Files: BEILSTEIN\*, BIOBUSINESS, CA, CAPLUS, CHEMCATS, CHEMLIST, CIN, IPA, PROMT, TOXCENTER, USAN, USPATFULL

(\*File contains numerically searchable property data)

Other Sources: EINECS\*\*

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

CM 1

CRN 77-92-9 CMF C6 H8 O7

CM 2

CRN 58-73-1 CMF C17 H21 N O

 $Ph_2CH-O-CH_2-CH_2-NMe_2$ 

CN

Carbowax 100

#### 15 REFERENCES IN FILE CA (1957 TO DATE) 15 REFERENCES IN FILE CAPLUS (1957 TO DATE) 137:103930 REFERENCE 1: REFERENCE 2: 137:11003 REFERENCE 135:262242 3: REFERENCE 132:255811 4: REFERENCE 129:265468 REFERENCE 129:127191 REFERENCE 7: 127:55916 REFERENCE 124:298541 REFERENCE 9: 121:42500 REFERENCE 10: 120:307174 ANSWER 3 OF 6 REGISTRY COPYRIGHT 2003 ACS RN **25322-68-3** REGISTRY Poly(oxy-1,2-ethanediy1), .alpha.-hydro-.omega.-hydroxy- (9CI) (CA INDEX CN NAME) OTHER NAMES: CN .alpha.,.omega.-Hydroxypoly(ethylene oxide) .alpha.-Hydro-.omega.-hydroxypoly(oxy-1,2-ethanediyl) CN .alpha.-Hydro-.omega.-hydroxypoly(oxyethylene) CN 1,2-Ethanediol, homopolymer CN 16600 CN CN 1660S 400DAB8 CN CN Alkox CN Alkox E 100 CN Alkox E 130 CN Alkox E 160 Alkox E 240 CN CN Alkox E 30 CN Alkox E 45 Alkox E 60 CN Alkox E 75 CN CN Alkox R 1000 CN Alkox R 15 CN Alkox R 150 CN Alkox R 400 . Alkox SR CN Antarox E 4000 CN CN Aquacide III CN Aquaffin CN Badimol CN BDH 301 Bradsyn PEG CN CN Breox 2000 Breox 20M CN Breox 4000 CN CN Breox 550 CN Breox PEG 300 CN CAFO 154 CN Carbowax

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CN
     Carbowax 1000
CN
     Carbowax 1350
     Carbowax 14000
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CN
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CN
     Carbowax 1540
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CN
     Carbowax 4600
ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
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     25104-58-9, 25609-81-8, 134919-43-0, 101677-86-5, 99264-61-6, 106186-24-7,
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     191743-71-2, 201163-43-1, 206357-86-0, 221638-71-7, 225502-44-3,
     270910-26-4, 307928-07-0, 356055-70-4, 391229-98-4
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CI
     PMS, COM
PCT
     Polyether
                  ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BIOBUSINESS, BIOSIS,
     STN Files:
LC
       BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CHEMSAFE, CIN, CSCHEM, CSNB, DDFU, DETHERM*,
       DIOGENES, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2,
       HSDB*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC,
       PDLCOM*, PIRA, PROMT, RTECS*, SPECINFO, TOXCENTER, TULSA, ULIDAT, USAN,
       USPAT2, USPATFULL, VETU, VTB
         (*File contains numerically searchable property data)
     Other Sources: DSL**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
```

69890 REFERENCES IN FILE CA (1957 TO DATE) 17762 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 70151 REFERENCES IN FILE CAPLUS (1957 TO DATE)

REFERENCE 1: 139:46031 REFERENCE 2: 139:45038 REFERENCE 3: 139;44671 REFERENCE 139:44654 REFERENCE 139:44568 5: 139:44208

6:

REFERENCE

7

CN

Butylenin

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REFERENCE
            7:
                139:43981
                139:43290
REFERENCE
            8:
REFERENCE
            9:
                139:43265
                139:42605
REFERENCE
           10:
    ANSWER 4 OF 6 REGISTRY COPYRIGHT 2003 ACS
L99
RN
     15687-27-1 REGISTRY
     Benzeneacetic acid, .alpha.-methyl-4-(2-methylpropyl)- (9CI)
                                                                      (CA INDEX
CN
     NAME)
OTHER NAMES:
CN
     (.+-.)-.alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid
CN
     (.+-.)-2-(p-Isobutylphenyl)propionic acid
CN
     (.+-.)-Ibuprofen
CN
     (.+-.)-Ibuprophen
     (4-Isobutylphenyl)-.alpha.-methylacetic acid
CN
CN
     (RS)-Ibuprofen
CN
     (S)-4-Isobutyl-.alpha.-methylphenylacetic acid
CN
     .alpha.-(4-Isobutylphenyl)propionic acid
CN
     .alpha.-Methyl-4-(2-methylpropyl)benzeneacetic acid
CN
     2-(4'-Isobutylphenyl)propionic acid
     2-(4-Isobutylphenyl)propanoic acid
CN
     2-(p-Isobutylphenyl)propionic acid
CN
     4-Isobutyl-.alpha.-methylphenylacetic acid
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CN
     4-Isobutylhydratropic acid
CN
     Act 3
     Adex 200
CN
CN
     Adran
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     Advil
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     Alaxan
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     Algofen
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CN
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CN
     Andran
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     Anflagen.
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     Antiflam
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     Apo-Ibuprofen
CN
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     Artofen
CN
     Artril
     Artril 300
CN
     Atril 300
CN
     Balkaprofen
CN
CN
     Betaprofen
CN
     Bloom
     Bluton
CN
CN
     Brofen
CN ·
     Brufanic
CN
     Brufen
CN
     Brufen 400
CN
     Brufen Retard
CN
     Bruflam
CN
     Brufort
CN
     Buburone.
CN
     Burana
CN
     Butacortelone
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Carol ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for DISPLAY FS 3D CONCORD 58560-75-1, 139466-08-3 DR C13 H18 O2 MF CI COM LC ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN\*, STN Files: BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DIOGENES, DIPPR\*, DRUGPAT, DRUGU, EMBASE, HSDB\*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, MSDS-OHS, NIOSHTIC, PHAR, PHARMASEARCH, PIRA, PROMT, RTECS\*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT, USAN, USPAT2, USPATFULL, VETU (\*File contains numerically searchable property data)

(\*\*Enter CHEMLIST File for up-to-date regulatory information)

DSL\*\*, EINECS\*\*, TSCA\*\*, WHO

Other Sources:

#### \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

6324 REFERENCES IN FILE CA (1957 TO DATE)
184 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
6364 REFERENCES IN FILE CAPLUS (1957 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 139:41987 REFERENCE 2: 139:41985 REFERENCE · 3: 139:41974 4: 139:41818 REFERENCE 139:41788 REFERENCE 5: 139:41655 REFERENCE 6: REFERENCE 7: 139:41618 REFERENCE 8: 139:41298

REFERENCE 9: 139:36906

REFERENCE 10: 139:30844

L99 ANSWER 5 OF 6 REGISTRY COPYRIGHT 2003 ACS

RN 147-24-0 REGISTRY

CN Ethanamine, 2-(diphenylmethoxy)-N,N-dimethyl-, hydrochloride (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Ethylamine, 2-(diphenylmethoxy)-N, N-dimethyl-, hydrochloride (8CI) OTHER NAMES:

```
CN
     .beta.-Dimethylaminoethyl benzhydryl ether hydrochloride
CN
     2-(Benzhydryloxy)-N, N-dimethylethylamine hydrochloride
CN
     2-Diphenylmethoxy-N, N-dimethylethylamine hydrochloride
CN
     Alledryl
CN
     Allergan
CN
     Allergina
CN
     Allergival
CN
     Amidryl
CN
     Bagodryl
CN
     Bax
CN
     Bena
CN
     Bena-Fedrin
CN
     Benadril hydrochloride
CN
     Benadryl
CN
     Benocten
CN
     Benodine
CN
     Benylin
CN
     Benzantin
CN
     Benzhydramine hydrochloride
     Benzhydrol .beta.-dimethylaminoethyl ether hydrochloride
CN
CN
     Denydryl
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     Dibondrin
CN
     Difenhydramine hydrochloride
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     Dimedrol
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     Dimedrol hydrochloride
CN
     Diphantine
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     Diphenhydramine hydrochloride
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     Fenylhist
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     Halbmond
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     Histacyl
CN
     Noctomin
CN
     Resmin
CN
     Restamin
CN
     S 8
CN
     Sedopretten
CN
     Sekundal D
CN
     Syntedril
CN
     Vena
CN
     Wehydryl
DR
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     COM
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LC
       BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST,
       CIN, CSCHEM, DETHERM*, DIOGENES, EMBASE, HODOC*, IFICDB, IFIPAT, IFIUDB,
       IPA, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PHARMASEARCH, PIRA, PROMT,
       RTECS*, SPECINFO, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
     (58 - 73 - 1)
CRN
Ph2CH-O-CH2-CH2-NMe2
```

**HCl** 

7.

1279 REFERENCES IN FILE CA (1957 TO DATE)

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10 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            1280 REFERENCES IN FILE CAPLUS (1957 TO DATE)
               8 REFERENCES IN FILE CAOLD (PRIOR TO 1967)
REFERENCE
            1: 139:41788
                139:26674
REFERENCE.
            2:
REFERENCE
            3:
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            7:
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REFERENCE
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REFERENCE
           10:
                138:343856
L99 ANSWER 6 OF 6 REGISTRY COPYRIGHT 2003 ACS
     58-73-1 REGISTRY
     Ethanamine, 2-(diphenylmethoxy)-N, N-dimethyl- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Ethylamine, 2-(diphenylmethoxy)-N, N-dimethyl- (7CI, 8CI)
OTHER NAMES:
     .alpha.-(2-Dimethylaminoethoxy)diphenylmethane
CN
     .beta.-Dimethylaminoethanol diphenylmethyl ether
CN
CN
     .beta.-Dimethylaminoethylbenzhydrylether
     2-(Benzhydryloxy)-N, N-dimethylethylamine
CN
     2-(Diphenylmethoxy)-N, N-dimethylethylamine
CN
CN
     Benzhydramine
CN
     Dimedrol base
CN
     Diphenhydramine
CN
     DPH
CN
CN
     N-[2-(Diphenylmethoxy)ethyl]-N, N-dimethylamine
CN
     O-Benzhydryldimethylaminoethanol
CN
     Probedryl
FS
     3D CONCORD
MF
     C17 H21 N O
CI
     COM
                  ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
LC
     STN Files:
       BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS,
       CHEMLIST, CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, HODOC*, HSDB*,
       IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, NAPRALERT, NIOSHTIC, PHAR,
       PHARMASEARCH, PROMT, RTECS*, SPECINFO, TOXCENTER, USAN, USPAT2,
       USPATFULL, VETU
         (*File contains numerically searchable property data)
                      EINECS**, NDSL**, TSCA**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
Ph2CH-O-CH2-CH2-NMe2
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3008 REFERENCES IN FILE.CA (1957 TO DATE)
62 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
3018 REFERENCES IN FILE CAPLUS (1957 TO DATE)
2 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

REFERENCE 1: 139:46005

REFERENCE 2: 139:41851

REFERENCE 3: 139:41788

REFERENCE 4: 139:26674

REFERENCE 5: 139:17578

REFERENCE 6: 139:17066

REFERENCE 7: 139:6872

REFERENCE 8: 138:396066

REFERENCE 9: 138:395915

REFERENCE 10: 138:390926

=> fil wpix

FILE 'WPIX' ENTERED AT 17:26:36 ON 16 JUL 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE LAST UPDATED: 16 JUL 2003 <20030716/UP>
MOST RECENT DERWENT UPDATE: 200345 <200345/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

- >>> NEW WEEKLY SDI FREQUENCY AVAILABLE --> see NEWS <<<
- >>> SLART (Simultaneous Left and Right Truncation) is now
   available in the /ABEX field. An additional search field
   /BIX is also provided which comprises both /BI and /ABEX <<</pre>
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<<
- >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES,
  SEE http://www.derwent.com/dwpi/updates/dwpicov/index.html <<<
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http://www.stn-international.de/training\_center/patents/stn\_guide.pdf <<</pre>

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- => d all abeq tech abex tot

L129 ANSWER 1 OF 3 WPIX (C) 2003 THOMSON DERWENT

AN **2002-590698** [63] WPIX

DNC **C2002-167152** 

TI Composition useful for treating pain-associated **sleep** disturbances comprises **ibuprofen** and **diphenhydramine**.

DC B05

IN BIANCO, J; COOK, G D; COOPER, S; DOYLE, G; FORT, J J; GIAMALVA, D H; KOCH,

T S; BIANCO, G; COOPER, S A; DOYLE, G A (BIAN-I) BIANCO J; (COOK-I) COOK G D; (COOP-I) COOPER S; (DOYL-I) DOYLE G; PA (FORT-I) FORT J J; (GIAM-I) GIAMALVA'D H; (KOCH-I) KOCH T S; (AMHP) WYETH CYC 100 75p WO 2002056877 A1 20020725 (200263)\* EN A61K031-00 PIRW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK. DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG UZ VN YU ZA ZM ZW US 2002177626 A1 20021128 (200281) A61K031-19 WO 2002056877 A1 WO 2002-US3855 20020118; US 2002177626 A1 Provisional US ADT 2001-262387P 20010119, US 2002-46727 20020117 PRAI US 2001-262387P 20010119; US 2002-46727 20020117 ICM A61K031-00; A61K031-19 A61K031-135 AB WO 200256877 A UPAB: 20021001 NOVELTY - A composition comprises ibuprofen (I) and diphenhydramine (II). DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for evaluating a compound or composition for the treatment of a pain-associated sleep disturbance in patients experiencing at least moderate or several levels of pain involving: (a) administering the compound or composition to the patient after oral surgery; (b) sending the patient to bed earlier than his usual bedtime; and (c) assessing the effectiveness of the compound or the composition. ACTIVITY - Analgesic; Sedative. Patients suffering from pain associated sleep disturbances were administered ibuprofen (400 mg) and diphenhydramine (50 mg) in a single soft gelatin capsule before bedtime. The symptoms of their pain associated sleep disturbances improved. They slept for a longer duration and also fell asleep faster than expected. Thus the composition was useful for treating pain associated sleep disturbances. MECHANISM OF ACTION - None given. USE - For treating pain-associated sleep disturbances affecting sleep duration (claimed) (preferably shortened sleep duration). The pain-associated sleep disturbances include difficulties in falling asleep and staying asleep. ADVANTAGE - The composition does not have negative interactions between (I) and (II) during formulation or when placed on shelves. The composition is stable chemically and physically. The combination of (I) and (II) yields a samnolent effect greater than that of (II) alone. (I) Alone is effective for only 4-6 hours while the composition is effective for longer periods of time. Dwg.0/29 FS CPI FΑ AB; DCN CPI: B10-B03B; B10-C02; B10-C04C; B11-C08; B12-K04; B12-M11B; MC B12-M11C; B12-M11K; B14-C01; B14-J01B1 TECH UPTX: 20021001 TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Composition: The composition comprises (I) (50-800, preferably 200 mg) and (II) in the form of diphenhydramine HCl (12.5-50, preferably 25 mg) or diphenhydramine citrate (19-38, preferably 38 mg). Preferred Method: The step (c) is performed by evaluating at least one of sleep latency, sum of pain relief and pain intensity differences over 0-3 hours (SPRID3), cumulative percent asleep at 60 minutes, ease of falling asleep, duration of sleep, global evaluation of sleep, median time to rescue medication, percent of patients requiring rescue medication, global assessment as a pain reliever, global

```
assessment as a sleep acid or pain relief intensity difference
     (PRID) 90 or 60 minutes (preferably sleep latency, sleep
     duration and SPRID3). The patient is sent to bed 1-6 (preferably at least
     1, 3 or upto 6) hours earlier than his usual bedtime.
ABEX
                    UPTX: 20021001
    ADMINISTRATION - Administration of (I) is 50-800 (preferably 200) mg. The
     dosage for diphenhydramine HCl is 12.5-50 (preferably
     25) mg and that for diphenhydramine citrate is 19-38
     (preferably 38) mg. The composition is a bilayer tablet or
     caplet or a soft gelatin capsule containing
    polyethylene glycol and is administered in one or two
     doses concomitantly. The second dose is optionally administered (all
     claimed). The composition is administered from 90 minutes before bedtime
     to immediately before bedtime (preferably 45-15 minutes especially 30
    minutes before bedtime).
     EXAMPLE - Diphenhydramine hydrochloride (40.050 kg)
     was dissolved in water (46.138 kg) and polyethylene
     glycol (molecular weight 600) (200.250 kg) was added to it.
     Polyethylene glycol (molecular weight 600) (326.808 kg)
     was also added to ibuprofen (320.400 kg) and mixed. Potassium
     hydroxide (41.011 kg) was dissolved in water (34.603 kg) and added to the
     ibuprofen dispersion and mixed under vacuum. The temperature was
     maintained at 75-125 degrees F. The diphenhydramine
    hydrochloride solution was added to the ibuprofen
     /potassium hydroxide/polyethylene glycol solution and
    mixed under vacuum. The solution was deaerated under vacuum for at least
     15 minutes and pressurized. The solution was encapsulated with
     in a gelatin shell and processed to completion. The capsule had
     the following composition (mg): ibuprofen (200),
     diphenhydramine HCl (25), DandC Red No. 33 (RTM)
     (0.0113), FDandC Blue No.1 (RTM) (0.0170), polyethylene
     glycol (molecular weight 600) (329), potassium hydroxide (25.6),
    purified water (50.4), ANDRISORB 85/70 (RTM) (93.4) and gelatin (164).
L129 ANSWER 2 OF 3 WPIX
                           (C) 2003 THOMSON DERWENT
    2001-290269 [30]
                       WPIX
ΑN
DNC
    C2001-088857
    Liquid softgel fill formulations comprising high concentrations of
TΙ
     ibuprofen in free form, and capsules containing them.
DC
    A14 A25 A96 B05 B07
IN
    GULLAPALLI, R P
PA
    (BANN-N) BANNER PHARMACAPS INC
CYC
PΙ
    WO 2001015688 A1 20010308 (200130) * EN
                                              20p
                                                     A61K031-19
       RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TZ UG ZW
        W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM
            DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC
            LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE
            SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW
    AU 2000070949 A
                     20010326 (200137)
                                                     A61K031-19
    US 6251426
                  B1 20010626 (200138)
                                                     A61K009-48
                                                                      <--
    EP 1207872
                  A1 20020529 (200243)
                                                     A61K031-19
                                         EN
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            RO SE SI
     JP 2003508434 W
                     20030304 (200319)
                                              21p
                                                     A61K031-192
    WO 2001015688 A1 WO 2000-US23908 20000901; AU 2000070949 A AU 2000-70949
ADT
     20000901; US 6251426 B1 US 1999-389003 19990902; EP 1207872 A1 EP
     2000-959667 20000901, WO 2000-US23908 20000901; JP 2003508434 W WO
     2000-US23908 20000901, JP 2001-519902 20000901
    AU 2000070949 A Based on WO 200115688; EP 1207872 A1 Based on WO
     200115688; JP 2003508434 W Based on WO 200115688
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PRAI US 1999-389003
                      19990902
     ICM A61K009-48; A61K031-19; A61K031-192
TC
         A61K009-06; A61K009-58; A61K009-64;
         A61K009-66; A61K045-00; A61K047-14; A61K047-22; A61K047-32;
         A61K047-34; A61P029-00
AΒ
    WO 200115688 A UPAB: 20010603
     NOVELTY - Stable liquid softgel fill formulations comprising high
     concentrations of ibuprofen in free form, polyethylene
     glycol and polyvinylpyrrolidone, and soft gelatin capsules
     containing them, are new.
          DETAILED DESCRIPTION - A liquid softgel fill formulation comprises
     (as wt.%):
          (a) greater than 30% ibuprofen (I) in free acid form in
     solution;
          (b) 30-60% polyethylene glycol (PEG);
          (c) 10-30% polyvinylpyrrolidone (PVP); and
          (d) optionally 1-10% surfactant to increase bioavailability of (I).
          INDEPENDENT CLAIMS are included for softgel capsules
     comprising a sheath enclosing a liquid fill as described above.
          USE - For administration of ibuprofen.
          ADVANTAGE - Use of a large amount of PVF permits a reduction in the
     amount of PEG required, reducing the esterification reaction
     between (I) and PEG.
     Dwg.0/0
FS
     CPI
FΑ
     AB; DCN
     CPI: A03-C01; A04-D05A; A05-H04; A12-V01; B04-C03C; B10-C04C; B12-M07;
MC
          B12-M11C
                    UPTX: 20010603
TECH
     TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Composition: The
     composition preferably comprises at least 35 wt.% (I); 30-50% PEG
     having average molecular weight (MW) 200-1,000; and 15-30% PVP having MW
     2,000-54,000. The surfactant is e.g. an ester of d-alpha-tocopheryl, a
     polyoxyethylene castor oil derivative, or polyglycolyzed
     glyceride.
     The composition may further comprise up to 6% of a nasal decongestant
     (e.g. pseudoephedrine hydrochloride), or up to 10% of an antihistaminic
     (e.g. diphenhydramine hydrochloride).
     The sheath of the softgel capsule comprises 45-75% gelatin;
     20-40% plasticizer (e.g. sorbitol, sorbitol special, and/or maltitol); and
     5-15% water. The total weight of the capsule fill is less than
     600 mg, and the weight of (I) is at least 200 mg.
ABEX
                    UPTX: 20010603
     EXAMPLE - A fill formulation was prepared comprising (as wt.%): (I) 35%;
     PEG 600 37%; Kollidon17 PF (RTM: PVP, MW 10,000) 22%; and Vitamin
     E TPGS 6%.
L129 ANSWER 3 OF 3 WPIX
                           (C) 2003 THOMSON DERWENT
AN
     1997-369376 [34]
                        WPIX
DNC
    C1997-118673
     Cold remedy soft capsule - comprises solution of
TI
     ibuprofen and second drug e.g. histamine antagonist.
DC
     A96 B05
PA
     (RPSE-N) RP SEALER KK; (SATO) SATO SEIYAKU KK
CYC
    1
                   A 19970617 (199734)*
                                              12p
                                                     A61K031-19
PΙ
     JP 09157162
                   B2 20030304 (200319)
                                              10p
                                                     A61K031-19
     JP 3382076
     JP 09157162 A JP 1995-322706 19951212; JP 3382076 B2 JP 1995-322706
ADT
     19951212
     JP 3382076 B2 Previous Publ. JP 09157162
FDT
                      19951212
PRAI JP 1995-322706
     ICM A61K031-19
         A61K009-08; A61K009-48; A61K031-52; A61K045-00;
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A61K045-06; A61K047-14; A61K047-34; A61P011-10; A61P011-14;
         A61P025-02; A61P025-04; A61P025-20; A61P025-26; A61P029-00;
         A61P043-00
    A61K031-19, A61K045:06
ICI
     JP 09157162 A UPAB: 19970820
AB
    A cold remedy soft capsule comprises a soln. of
     ibuprofen and one or more second drugs from histamine antagonist,
     antitussive agent, expectorant, sympathetic nerve stimulant and central
     stimulant in a base contg. a surfactant from polyoxyethylene
     (POE) sorbitan fatty acid ester, POE hardened castor oil and polyglycerol
     fatty acid ester and water, and no hydroxide ions.
          The central stimulant is anhydrous caffeine, caffeine or sodium
     benzoate caffeine.
          The surfactant has HLB of 13-18.
          POE (20) sorbitan monooleate (1030.5g) at about 60 deg.C was added
     with ibuprofen (225g), diphenhydramine
    hydrochloride (37.5g), dihydrocodeine phosphate (12b), noscapine
     hydrochloride (24mg), dl-methylephedrine hydrochloride (30g), guaifenesin
     (125g) and anhydrous caffeine (15g). The mixt. was vigorously stirred
     using a homogenizer, cooled to about 40 deg.C, added with purified water
     (106q), deaerated under vacuum and filtered through a 100 mesh. The
     filtrate (each 535mg) was filled in a gelatin film [comprising gelatin
     (100g) and glycerol (30g)] to give 3,000 soft capsules.
          ADVANTAGE - Absorption of the drugs is excellent, ibuprofen
     is stable in the capsule, and the capsule need not be
     measured and is handy to carry.
     Dwq.0/0
FS
     CPI
FΑ
     CPI: A10-E07; A12-V01; B04-A04; B04-A06; B04-A10; B04-N02; B05-A01B;
MC
          B10-B03B; B10-C04C; B12-M09; B12-M11C; B14-C04; B14-L09
=> d his
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                S 15687-27-1/REG#
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                $ 58-73-1/REG#
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L2
L3
              1 S 58-73-1
              1 S 25322-68-3
L4
            137 S C13H18O2/MF AND 46.150.18/RID AND BENZENEACETIC
L5
              3 S L5 AND IBUPROFEN
L6
              3 S L2, L6
L7
rac{1}{8}
            209 S 58-73-1/CRN
                SEL RN L7
            313 S E1-E3/CRN
L9
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4 S L8 AND L9
L10
L11
             1 S L10 AND 2/NC
L12
           110 S L8 NOT (MXS OR IDS OR PMS)/CI
L13
            29 S L12 NOT (UNSPECIFIED OR COMPD OR WITH)
L14
             26 S L13 NOT SALT
L15
             23 S L14 NOT (CONJUGATE OR CR/ELS)
L16
             3 S L13 NOT L14
             26 S L15,L16
L17
L18
             1 S L17 AND CLH
L19
             2 S L17 AND C6H8O7
L20
             3 S L18, L19
L21
             23 S L17 NOT L20
             44 S L9 NOT ((MXS OR IDS OR PMS)/CI OR UNSPECIFIED OR COMPD OR WI
L22
L23
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L25
           6751 S L7 OR L23
L26
           7215 S IBUPROFEN
L27
            27 S DEXIBUPROFEN
L28
            140 S 2 4 ISOBUTYLPHENYL PROPIONIC ACID
L29
            39 S 2 P ISOBUTYLPHENYL PROPIONIC ACID
L30
           8206 S L25-L29
L31
          1294 S L20 OR L21
L32
           4208 S DIPHENHYDRAMINE
L33
            16 S DIPHENHYDRAMINE CITRATE
L34
           527 S DIPHENHYDRAMINE HYDROCHLORIDE
L35
           459 S DIPHENHYDRAMINE (A) HCL
L36
           535 S DIMEDROL
L37
           5035 S L31-L36
L38
            227 S L30 AND L37
L39
             33 S L38 AND L4
             33 S L38 AND (PEG OR POLYETHYLENEGLYCOL OR POLYETHYLENEOXIDE OR PO
L40
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L41
             40 S L39, L40
L42
             23 S L38 AND POLYOXYALKYLENE
L43
             42 S L42, L43
L44
L45
              1 S L44 AND SLEEP?
                E SLEEP/CT
           7983 S E3-E10
L46
                E E3+ALL
           8142 S E2+NT
L47
           1580 S E3, E4/BI
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             1 S L44 AND L46-L48
L50
             1 S L38 AND L46-L48
             1 S L38 AND SLEEP?
L51
                E INSOMNIA/CT
L52
            570 S E3, E4
               E E3+ALL
L53
              0 S L38 AND L52
L54
              0 S L38 AND INSOMN?
L55
              1 S L49-L51
L56
           3046 S L3
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L58
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L63
L64
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                E PAIN/CT
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E E3+ALL

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L72
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L75
                E COOPER STEPHEN/AU
L76
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                E COOK G/AU
L77
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                E COOK GRAHAM/AU
L78
              9 S E3-E5
                E KOCH T/AU
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L82
             21 S E3-E7
              1 S E19
L83
                E FORT J/AU
L84
             52 S E3, E8-E11
                E DOYLE G/AU
L85
             59 S E3-E11
L86
              7 S E38-E40
L87
              1 S L74-L86 AND L62
L88
              2 S WYETH?/PA,CS AND L62
             2 S L61, L87, L88
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L94
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L95
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L96
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          32310 S (POLYETHYLENE()(GLYCOL OR OXIDE) OR POLYOXY ETHYLENE OR POLY
L115
            35 S L111 AND L112-L115
L116 .
            16 S L116 AND (?CAPSUL? OR ?CAPLET? OR ?TABLET?)/BIX
L117
            10 S L116 AND (R031 OR R033 OR R034 OR R038)/M0,M1,M2,M3,M4,M5,M6
L118
            14 S L116 AND (B12-M11 OR B12-M11A OR B12-M11B OR B12-M11C OR B12-
L119
            0 S L116 AND (C12-M11 OR C12-M11A OR C12-M11B OR C12-M11C OR C12-
L120
            17 S L117-L119
L121
            18 S L116 NOT L121
L122
L123
            27 S L116 AND A61K009/IC, ICM, ICS, ICA, ICI
L124
            15 S L123 AND L121
            17 S L121, L124
L125
            12 S L123 NOT L125
L126
            1 S L125 AND SLEEP?/BIX
L127
            17 S L125, L127 AND M782/M0, M1, M2, M3, M4, M5, M6
L128
                SEL DN AN 5 8 11
L129
              3 S L128 AND E1-E6
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